

## Navigating the ENCODE Encyclopedia: Exploring Candidate Regulatory Elements, Linked Genes, and Genetic Variation with SCREEN



Jill Moore, Michael Purcaro, and Zhiping Weng University of Massachusetts Medical School

# Part II – Overview of The Registry of Candidate Regulatory Elements

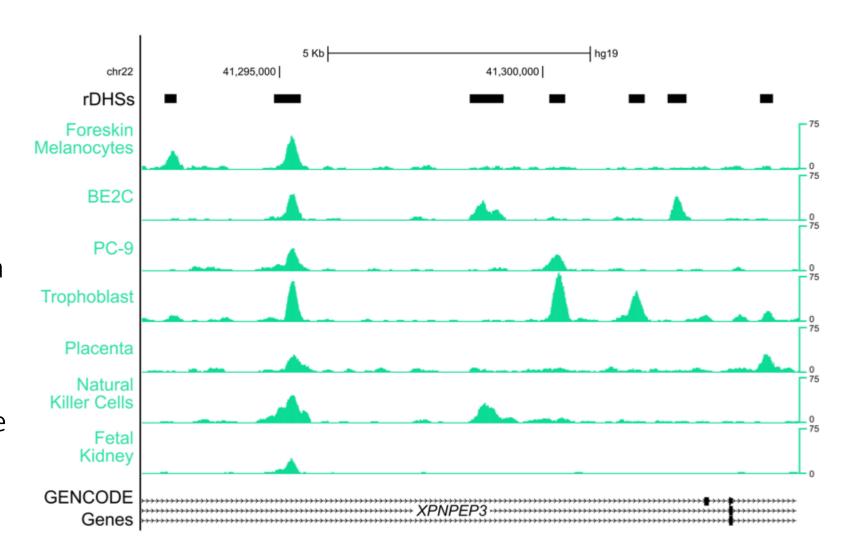
Jill Moore ENCODE DAC Project Manager

#### ENCODE Registry of <u>candidate Regulatory Elements</u> (cREs)

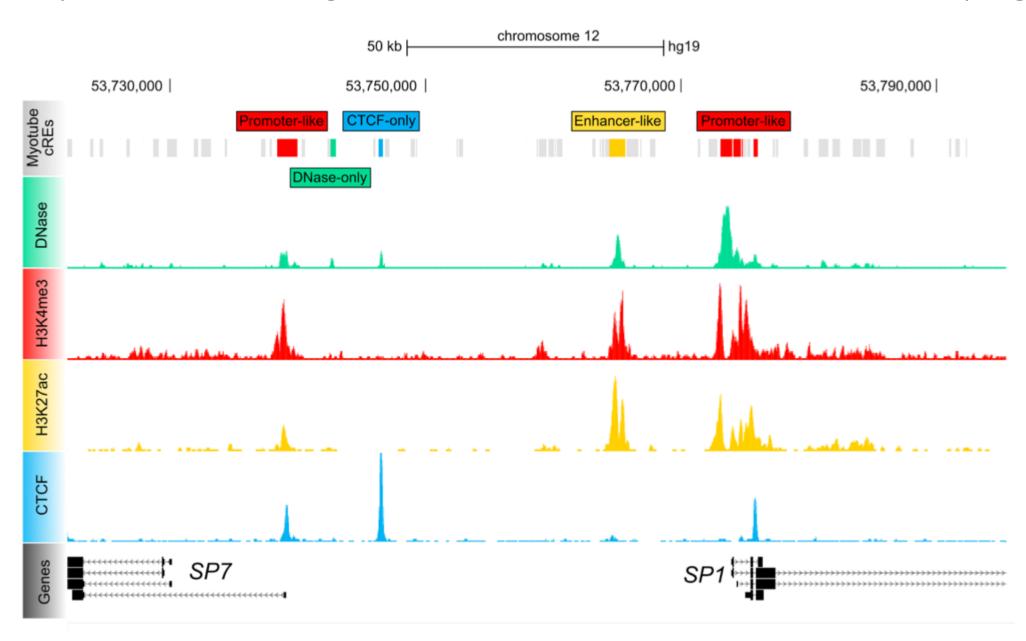
- Genomic regions that may function as regulatory elements including promoters, enhancers, and insulators
- Incorporate data from hundreds of cell and tissue types
- Creating registries in both human and mouse

#### Step 1 – Creating representative DHSs (rDHSs)

- We generate DHS clusters using DHSs across over 400 cell types
- For each cluster we select a representative DHS (rDHS)
- We iteratively run this procedure until all DHSs are represented



#### Step 2 – Annotating rDHSs with Histone & CTCF ChIP-seq Signal

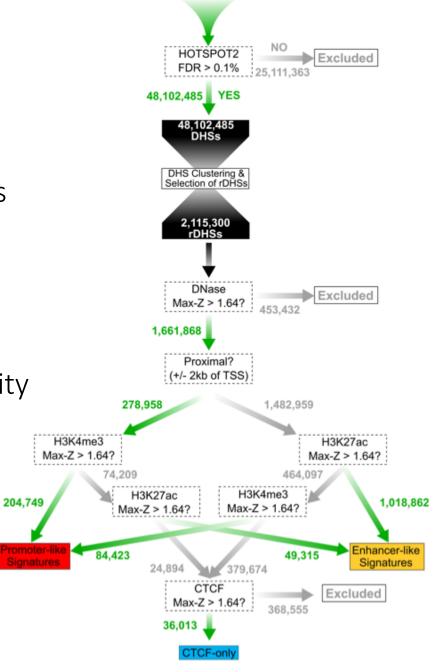


#### Two Types of Classification Schemes:

- 1. Cell type agnostic:
  - Independent of cell type
  - Use maximum ChIP-seq signal across all cell types and distance from TSS

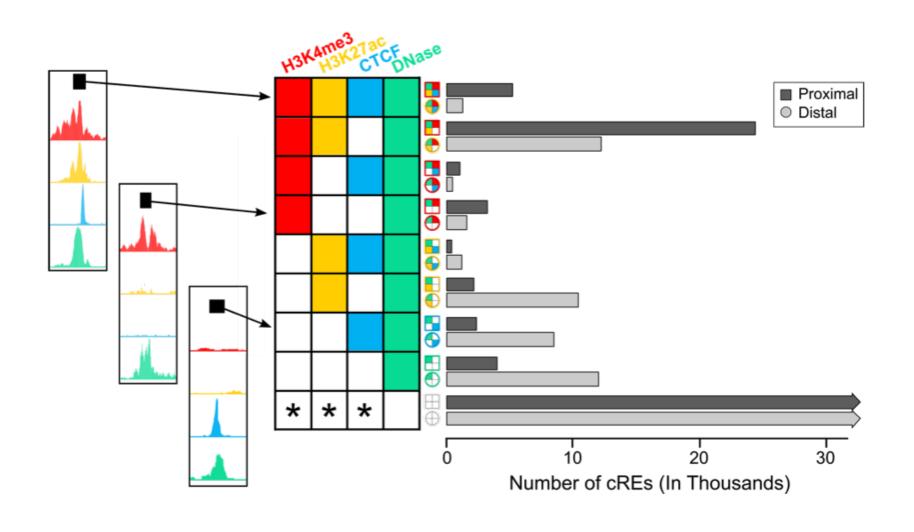
2. Cell type specific:

 Use cell-type specific signal to annotate the activity of each cRE in over 600 cell types



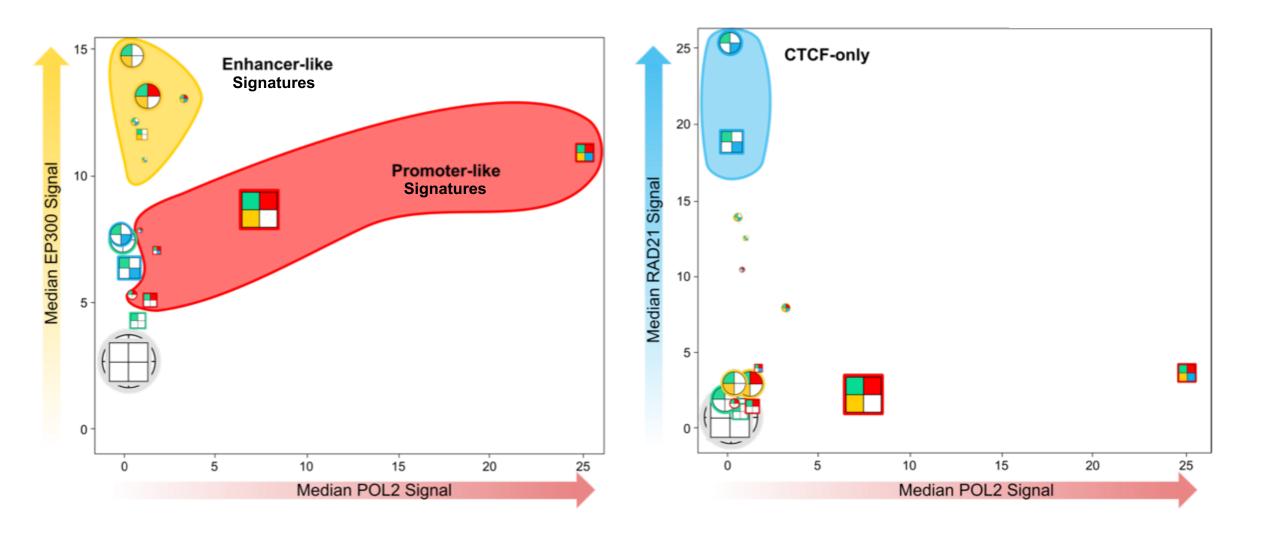
73,213,848

#### Cell Type Specific Classification

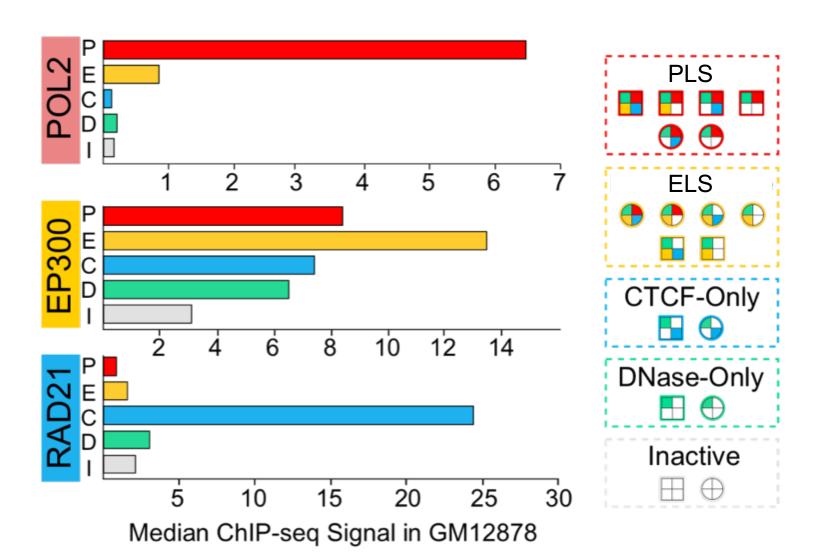


- H3K4me3
- H3K27ac
- CTCF
- distance to the nearest TSS

#### Validation using Pol II, EP300, and RAD21 TF ChIP-seq Signal



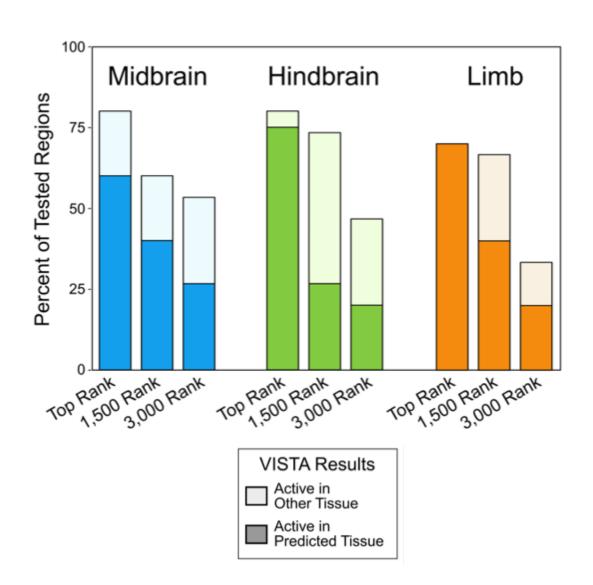
#### Simplifying Cell Type Specific States into 5 Groups

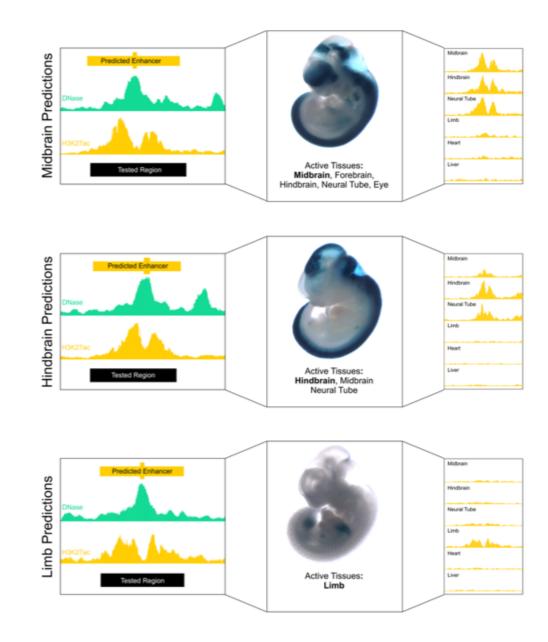


rDHSs are classified into five groups:

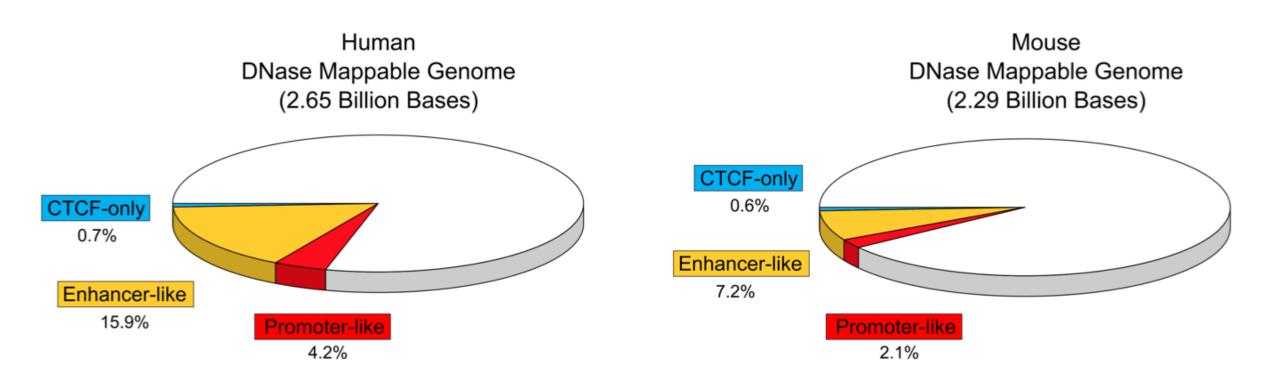
- Promoter-like signatures
- Enhancer-like signatures
- CTCF-only
- DNase-only
- Inactive

#### Validation of 151 ELS cREs using transgenic assays

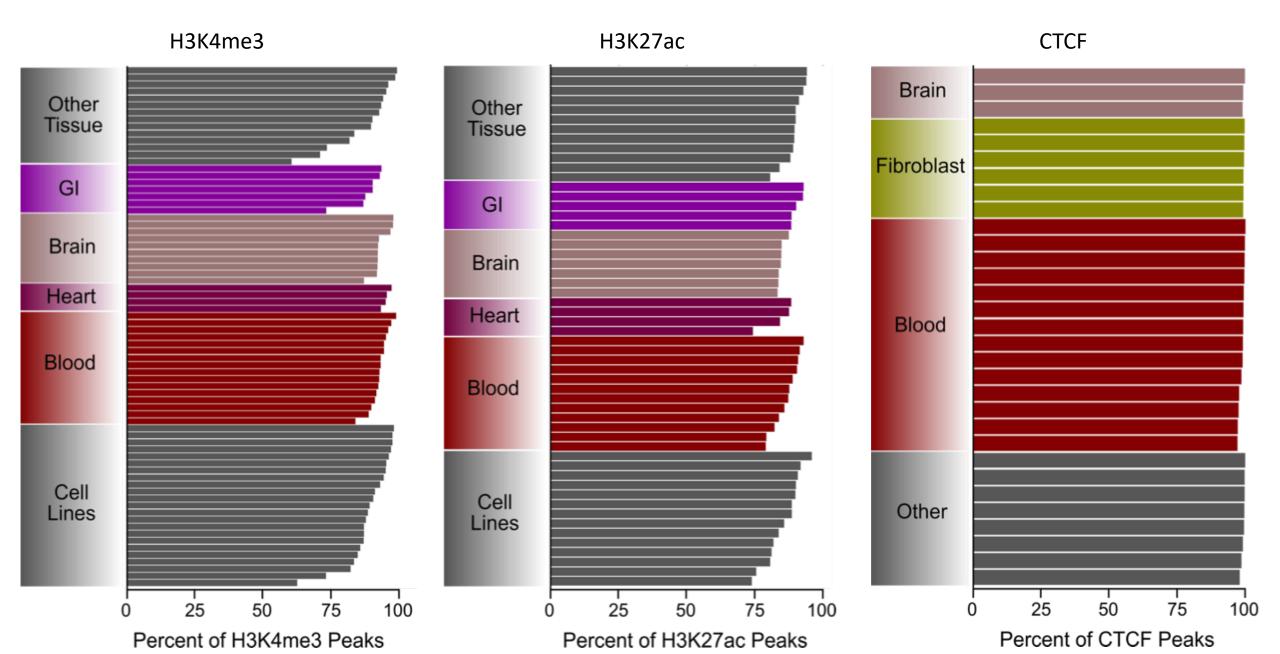




# In total, we curated 1.3 million cREs in human and 431 thousand cREs in mouse.



#### Coverage of the Human cRE Registry in New Cell Types





### Disclosure for:

ASHG Interactive Workshop: Navigating the ENCODE Encyclopedia: Exploring Candidate Regulatory Elements, Linked Genes, and Genetic Variation with SCREEN

No Relevant Conflicts to Disclose:

Jill Moore

**Zhiping Weng** 

Michael Purcaro